(19) World Intellectual Property Organization International Bureau



(43) International Publication Date 8 September 2000 (08,09,2000)

English

(10) International Publication Number WO 00/51633 A3

- 39/116, 39/295, A61P 31/16
- (21) International Application Number: PCT/CA00/00207
- (22) International Filing Date: 29 February 2000 (29.02.2000)

(25) Filing Language:

- (26) Publication Language: English
- (30) Priority Data: 09/261.182 3 March 1999 (03.03.1999) US
- (71) Applicant (for all designated States except US): CON-NAUGHT LABORATORIES LIMITED [CA/CA]: 1755 Steeles Avenue West, Toronto, Ontario M2R 3T4 (CA).
- (72) Inventors: and
- (75) Inventors/Applicants (for US only): LOOSMORE, Sheena, M. [CA/CA]; 70 Crawford Rose Drive, Aurora, Ontario L4G 4R4 (CA), YANG, Yan-Ping [CA/CA]; Apt. 709, 120 Torresdale Avenue, Willowdale, Ontario M2R 3N7 (CA), KLEIN, Michel, H. [CA/CA]: 16 Munro Boulevard, Willowdale, Ontario M2P 1B9 (CA).

- (51) International Patent Classification?: A61K 39/102. (74) Agent: STEWART, Michael, L.: Sim & McBurney, 6th Floor, 330 University Avenue, Toronto, Ontario M5G 1R7 (CA).
 - (81) Designated States (national): AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, IP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT. RO, RU, SD, SE, SG, SL, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW,
 - (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU. MC, NL, PT, SE). OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Published:

- --- With international search report.
- (88) Date of publication of the international search report: 25 January 2001

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: MULTI-COMPONENT VACCINE COMPRISING AT LEAST THREE ANTIGENS TO PROTECT AGAINST DIS-EASE CAUSED BY HAEMOPHILUS INFLUENZAE

(57) Abstract: A multi-component immunogenic composition confers protection on an immunized host against infection caused by Haemophilus influenzae. Such composition comprises at least three different antigens of Haemophilus influenzae, two of which are adhesins. High molecular weight (HMW) proteins and Haemophilus influenzae adhesin (Hig) proteins of non-typeable Haemophilus influenzae comprise the adhesin components while the other amagen is a non-proteolytic analog of Hia47 protein. Each component does not impair the immunogenicity of the others. The Haemophilus vaccine may be combined with DTP component vaccines. which may contain inactivated poliovirus, including type 1, type 2 and/or type 3, and/or a conjugate of a capsular polysaccharide of Haemophilus influenzas and tetanus or diphtheria toxoid, including PRP-T, to provide a multi-valent component vaccine without impairment of the immunogenic properties of the other antigens.

INTERNATIONAL SEARCH REPORT

interr nai Application No PCT/CA 00/00207

A CLASSIFICATION OF SUBJECT MATTER
1PC 7 A61K39/102 A61K39/116 A61K39/295 A61P31/16 According to International Patent Classification (IPC) or to both national classification and IPC B REIDS SEARCHED Minimum documentation seatened (classification system followed by classification symbols) A61K Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted coving the international search (name of data base and, where practical, search terms used) EPO-Internal, BIOSIS, PAJ, WPI Data, MEDLINE C. DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. BARENKAMP STEPHEN J ET AL: 1-4.7-9 "Identification of a second family of cited high-molecular-weight adhesion proteins expressed by non-typable Haemophilus influenzae. MOLECULAR MICROBIOLOGY. vol. 19, no. 6, 1996, pages 1215-1223, XP000946619 ISSN: 0950-382X cited in the application page 1215, column 2, paragraph 1 page 1220, column 2 -page 1221, column 1, paragraph 1 -/---X Further documents are listed in the continuation of box C. X Patent family members are listed in annex. Special categories of cited documents: *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention. "A" document defining the general state of the art which is not considered to be of particular relevance. "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken along Sing date *L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of anothe citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention carnot be considered to involve an inventive step when the document is combined with one or more other, such docu-"O" document reterring to an onal disclosure, use, exhibition or other means ments, such combination being obvious to a person skilled "P" document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international awards report 10 October 2000 17/10/2000 Name and mailing address of the ISA Authorized officer European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Pijswilk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016 Noë. V

INTERNATIONAL SEARCH REPORT

Intern nel Application No PCT/CA 00/00207

A S BARENKAMP: "Immunization with high-molecular weight adhesion proteins of nontypeable H. influenzae modifies experimental otitis media in chincillas" INFECTION AND IMMUNITY, AMERICAN SOCIETY OF MICKOBIOLOGY, WASHINGTON, DC, US,	1-3,7-9, 18.20,21
A S BARENKAMP: "Immunization with high-molecular weight adhesion proteins of nontypeable H. influenzae modifies experimental otitis media in chincillas" INFECTION AND IMMUNITY, AMERICAN SOCIETY OF MICROBIOLOGY, WASHINGTON, DC, US,	1-3,7-9,
high-molecular weight adhesion proteins of nontypeable H. influenzae modifies experimental otitis media in chincillas" INFECTION AND IMMUNITY, AMERICAN SOCIETY OF MICROBIOLOGY, WASHINGTON, DC, US,	
vol. 64, no. 4, 1996, pages 1246-1251, XP002142962 ISSN: 0019-9567 cited in the application abstract page 1250	10,20,21
	1-3,7-9, 18-21
US 5 869 302 A (LOOSMORE SHEENA M ET AL) 9 February 1999 (1999-02-09) the whole document	1,5.6, 8-15
E WO 00 35477 A (LOOSMORE SHEENA M ;CONNAUGHT LAB (CA); YANG YAN PING (CA); KLEIN M) 22 June 2000 (2000-06-22) cited in the application the whole document	1-27

INTERNATIONAL SEARCH REPORT

amation on patent family members

Interr nai Application No PCT/CA 00/00207

	ent document n search repor	t	Publication date		Patent family member(s)		Publication date
WO !	9421290	A	29-09-1994	AU	696207	B	03-09-1998
				AU	6400594		11-10-1994
				BR		A	30-01-1996
				EP		A	03-01-1996
				JP	11501003	T	26-01-1999
				US	5869065	À	09-02-1999
US :	5869302	A	09-02-1999	US	5939297	A	17-08-1999
				US	5506139	A	09-04-1996
			AU	687619	В	26-02-1998	
				AU	3337695	Α	22-02-1996
			BR	9506272	Α	12-08-1997	
			CA	2171611	A	08-02-1996	
			WO	9603506	A	08-02-1996	
			ΕP		A	04-09-1996	
				NZ		A	24-10-1997
				US		A	15-02-2000
				US		A	01-02-2000
			US	6114125	A	05-09-2000	
				CN		A	20-11-1996
				US	5665353		09-09-1997
				US	5935573		10-08-1999
				US	5656436		12-08-1997
				US	5981503		09-11-1999
				US	5962430	A	05-10-1999
MU	0035477	A	22-06-2000	AU	1543900	٨	03-07-2000